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		PPR	Abrey et al., Combination clymphoma," (abstract) Proc. Ar	hemothera	apy in primary centra					
		QQR	Abrey <i>et al.</i> , Long-term surv 859-63 (1998)	vival in prir	mary CNS lymphoma	ı, <u>J. Clin. Oncol.</u> 16:				<u> </u>
		RRR	Armitage <i>et al.,</i> Molecular a CD40, <u>Nature</u> 357:80-82 (1992		cal characterization o	of a murine ligand for				_
-		SSR	Behr <i>et al.</i> , Low-versus high CD22 or chimeric anti-CD20 ar malignancies, <u>Clin. Cancer Re</u>	ntibodies ii	n a broad spectrum o					
		TTR	Bejcek et al., Development chain antibody fragments (scFv 55: 2346-51 (1995)							
		UUR	Benoit et al., Increased inhi following ligation of CD40, and immunoglobulin, Immunopharn	l either CD	19, CD20, CD95 or s				,,,,	
		VVR	Berinstein <i>et al.</i> , Association and anti-tumor response in the Hodgkin's lymphoma, Ann. On	treatment	of recurrent low-gra-					
		WWR	Blay et al., High-dose meth- lymphomas: Analysis of surviva series, <u>J. Clin. Oncol.</u> 16: 864-8	otrexate for	or the treatment of prince neurologic toxicity in					
		XXR	Bolognesi <i>et al.</i> , Evaluation inactivating proteins and an an <i>in vivo</i> studies, Br. J. Haemato	of immun nti-CD22 m	otoxins containing si onoclonal antibody (
		YYR	Capon <i>et al.,</i> Designing CD 525-531 (1989)			erapy, <u>Nature</u> 337:		-		
		ZZR	Chamberlain et al., Priman adjuvant chemotherapyt, <u>J. Ne</u>							
		AAAR	Cheng <i>et al.</i> , Systemic che immunodeficiency syndrome-re 82: 1946-51 (1998)				:			
		BBBR	Clodi <i>et al.</i> , Unbalanced ex Brit. J. Haematol. 103: 217-9 (of Fas and CD40 in n	nantle cell lymphoma,				
		CCCR	Clynes et al., Fc receptors melanoma, PNAS (USA) 95:65			tive immunity to				
		DDDR	Coiffier et al., Rituximab (al patients with relapsing or refractudy, Blood 92: 1927-1932 (19	ctory aggre						
		EEER	Czuczman <i>et al.</i> , Treatmen the combinations of chimeric a chemotherapy, <u>J. Clin. Oncol.</u>	anti-CD20 r	monoclonal antibody					
	 -	FFFR	DeAngelis et al., "Primary (PRINCIPLES & PRACTICE OF ONCO							
		GGGR	Deguchi <i>et al.</i> , Retention o following conjugation of a bloop poly(ethylene glycol) linker, <u>Bio</u>	d-brain ba	rrier drug delivery ve	ctor via an extended				
		HHHR	Endo, <u>Gan To Kagaku Rye</u>	oho 26: 74	1-748 (1999)			_		†

	, W	1IIR	Fine et al mary central nervous system lymphoma, Annual ntern. Med. 119: 1093-1104 (1993)	-		ECA	
SE	1 P	703 48	Flavell et al., Therapy of human B-cell lymphoma bearing SCID mice is more effective with anti-CD19- and anti-CD38-saporin immunotoxins used in combination than with either immunotoxin used alone, Int. J. Cancer 62: 337-44 (1995)	TECH OF	VIE)	0,20	23
TENT & T	RADEN	KKKR	Freilich <i>et al.</i> , Chemotherapy without radiation therapy as initial. Treatment for primary CNS lymphoma in older patients, <u>Neurology</u> 46: 435-439 (1996)			1600/23	00
		LLLR	Funakoshi <i>et al.</i> , Inhibition of human B-cell lymphoma growth by CD40 stimulation, <u>Blood</u> 83: 2787-2794 (1994)	-			
		MMMR	Funakoshi <i>et al.</i> , Differential in vitro and in vivo antitumor effects mediated by anti-CD40 and anti-CD20 monoclonal antibodies against human B-cell lymphomas, <u>J. Immunother. Emphasis Tumor Immunol.</u> 19: 93-101 (1996)				
		NNNR	Ghetie <i>et al.</i> , Anti-CD19 antibodies inhibit the function of the P-gp pump in multigrug-resistant B lymphoma cells, <u>Clin. Cancer Res.</u> 5: 3920-7 (1999)				
		OOOR	Green et al., Evidence for a continued requirement for CD40/CD40 ligand (CD154) interactions in the progression of LP-BM5 retrovirus-induced murine AIDS, Virology 241: 260-268 (1998)				
		PPPR	Gruss <i>et al.</i> , CD40/CD40 ligand interactions in normal, reactive and malignant lympho-hematopoietic tissues, <u>Leuk. Lymphoma</u> 24: 393-422 (1997)				
		QQQR	Hekman et al., Initial experience with treatment of human B cell lymphoma with anti-CD19 monoclonal antibody Cancer Immunol. Immunother. 32:364-372 (1991)				
		RRRR	Herrlinger <i>et al.</i> , Intrathecal therapy of leptomeningeal CEM T-cell lymphoma in nude rats with anti-CD7 ricin toxin A chain immunotoxin, <u>J. Neurooncol.</u> 40: 1-9 (1998)				
		SSSR	Hollenbaugh <i>et al.</i> , The human T cell antigen gp39, a member of the TNF gene family, is a ligand for the CD40 receptor: expression of a soluble form of gp39 with B cell co-stimulatory activity, <u>EMBO J.</u> 11:4313-4321 (1992)				
		TTTR	Hollinger et al., "Diabodies": small bivalent and bispecific antibody fragments, Proc. Nad. Acad. Sci. USA, 90:64446448 (1993).				
		UUUR	Huwyler <i>et al.</i> , Brain drug delivery of small molecules using immunoliposomes, Proc. Nat'l Acad. Sci. USA 93: 14164-14169 (1996)				
		VVVR	Illidge et al., The importance of antibody-specificity in determining successful radioimmunotherapy of B-cell lymphoma, <u>Blood</u> 94: 233-43 (1999)				
		WWWR	Janeway, Immunotherapy by peptides, Nature, 341: 482 (1989)				
		XXXR	Johnson et al., Isolated follicular lymphoma cells are resistant to apoptosis and can be grown in vitro in the CD40/stromal cell system, <u>Blood</u> 82: 1848-1857 (1993)				
		YYYR	Kiesel et al., Removal of cells from a malignant B-cell line from bone marrow with immunomagnetic beads and with complement and immunoglobulin switch variant mediated cytolysis, <i>Leukemia Research 11</i> , 12: 1119 (1987)				
		ZZZR	Kramer et al., Tc-99m LL-2 Fab' monoclonal antibody imaging in acquired immune deficiency syndrome-related lymphoma, <u>Cancer</u> 80: 2469-2477 (1997)				
		AAAAR	Kroll <i>et al.</i> , Outwitting the blood-brain barrier for therapeutic purposes: Osmotic opening and other means, Neurosurgery 42: 1083-99 (1998)				
		BBBBR	Ledbetter <i>et al.</i> , Agonistic and antagonistic properties of CD40 mAb G28-5 are dependent on binding valency, <u>Circ. Shock</u> 44: 67-72 (1994)				

					. 4	\$ _	
		CCCCR	Led'ermal 1., Identification of a novel surface protein on vated CD4+ T cells that induces contact-dependent B cell differentiation (Help). Exp. Med. 175: 1091-1101 (1992)	Ec.	OC	'CC	
P 2 6	l	DDDDR	Leget <i>et al.</i> , Use of rituximab, the new FDA-approved antibody, <u>Curr. Opin.</u> Oncol. 10: 548-551 (1998)	CHC	WZ	910	70;
& TRADE	مر	EEER	Lesser et al., The chemotherapy of adult primary brain tumor, Cancer Treat. Rev. 19: 261-281 (1993)			.000/2	300
		FFFFR	Li et al., Pharmacokinetics and biodistribution of radioimmunoconjugates of anti-CD19 antibody and single-chain Fv for treatment of human B-cell malignancy, Cancer Immunol. Immunother. 47: 121-30 (1998)				
		GGGGR	Lieberman <i>et al.</i> , Convection-enhanced distribution of large molecules in gray matter during interstitial drug infusion, <u>J. Neurosurg.</u> 82: 1021-1029 (1995)				
-		ннннк	Linsley et al., Binding of the B cell activation antigen B7 to CD28 costimulates T cell proliferation and interleukin 2 mRNA accumulation, <u>J. Exp. Med.</u> 1783: 721-730 (1991)				
		IIIIR	Maloney et al., IDEC-C2B8 (Rituximab) anti-CD20 monoclonal antibody therapy in patients with relapsed low-grade non-Hodgkin's lymphoma, <u>Blood</u> 90: 2188-2195 (1997)				
		JJJJR	Mansfield <i>et al.</i> , Characterization of RFB4-pseudomonas exotoxin A immunotoxins targeted to CD22 on B-cell malignancies, <u>Bioconjug. Chem.</u> 7: 557-63 (1996)				
		KKKKR	Mason <i>et al.</i> , ¹¹¹ Indium-diethylenetriamine pentaacetic acid cerebrospinal fluid flow studies predict distributin of intrathecally administered chemotherapy and outcome in patients with leptomeningeal metastases, <u>Neurology</u> 50: 438-444 (1998)				
		LLLLR	McLaughlin <i>et al.</i> , Rituximab chimeric anti-CD20 monoclonal antibody therapy for relapsed indolent lymphoma: Half of patients respond to a four-dose treatment program, J. Clin. Oncol. 16: 2825-2833 (1998)	·			
		MMMMR	McLaughlin <i>et al.</i> , Clinical status and opitimal use of rituximab for B-cell lymphomas, Oncology (Huntingt) 12: 1763-1777 (1998)				T
+		NNNNR	Monjour <i>et al.</i> , Lymphomes malins non Hodgkiniens primitifs du système nerveux central, Rev. Neurol. (Paris) 148: 589-600 (1992)				+
di	2	0000R	Morrison <i>et al.</i> , Genetically engineered antibody molecules, <u>Adv. Immunol.</u> 44: 65-92 (1988)				
7		PPPPR	Morrison et al., Chimeric human antibody molecules: Mouse antigen-binding domains with human constant region domains, <i>Proc. Natl. Acad. Sci.</i> USA, 81:6851-6855 (1984)				
		QQQQR	Murphy et al., Antibodies to CD40 prevent Epstein-Barr virus-mediated human B-cell lymphomagenesis in severe combined immune deficient mice given human peripheral blood lymphocytes, Blood 86: 1946-1953 (1995)				
	-	RRRRR	Nguyen et al., IDEC-C2B8 anti-CD20 (Rituximab) immunotherapy in patients with low-grade non-Hodgkin's lymphoma and lymphoproliferative disorders: evaluation of response on 48 patients, <u>Eur. J. Haematol.</u> 62: 76-82 (1999)	•			
		SSSSR	Offner et al., T Cell receptor peptid the rapy triggers autoregulation of experimental encephalomyelitis, Science 251: 430-432 (1991)			-	1
		TTTTR	O'Neill <i>et al.</i> , Primary central nervous system non-Hodgkin's lymphoma: Survival advantages with combined initial therapy?, Int'l J. Radiation Oncol. Biol. Phys. 33: 663-673 (1995)				+-

					۵.	
60	UUUUR	Orlówski Hodgkin's disease with leptomeningeal invo		0		8
	VVVVR	Padlan, A possible procedule for reducing the immunogenicity of antibody variable domains while preserving their ligand-binding properties, <u>Molec. Immun.</u> 28: 489-498 (1991)	TECH (ENZ	70,	200
	WWWWR	Padlan, Anatomy of the antibody molecule, Molec. Immun. 31: 169-217 (1994)			1600	200
	XXXXR	Partridge <i>et al.</i> , Combined use of carboxyl-directed protein pegylation and vector-mediated blood-brain barrier drug delivery system optimizes brain uptake of brain-derived neurotrophic factor following intravenous administration, <u>Pharm.</u> <u>Res.</u> 15: 576-82 (1998)			, , , , , , , , , , , , , , , , , , ,	
	YYYYR	Pastan <i>et al.</i> , Intrathecal administration of single-chain immunotoxin, LMB-7 [B3(Fv)-PE38], produces cures of carcinomatous meningitis in a rat model, <u>Proc. Nat'l Acad. Sci. USA</u> 92: 2765-2769 (1995)				
-	ZZZZR	Perez-Jaffe <i>et al.</i> , Cerebral spinal fluid involvement by Hodgkin's disease diagnosed by CSF cytology and immunocytochemistry, <u>Diagn. Cytopathol.</u> 20:219-223 (1999)			••	
	AAAAAR	Piro <i>et al.</i> , Extended Rituximab (anti-CD20 monoclonal antibody) therapy for relapsed or refractory low-grade or follicular non-Hodgkin's lymphoma, <u>Ann. Oncol.</u> 10: 655-61 (1999)			· -	
	BBBBBR	Press <i>et al.</i> , Monoclonal antibody 1F2 (Anti-CD20) serotherapy of human B cell lymphomas, <u>Blood</u> 69: 584-591 (1987)				
	CCCCCR	Ravetch and Kinet, Fc receptors, Annu. Rev. Immunol 9:457-92 (1991)	1 24			
	DDDDDR	Reni <i>et al.</i> , Therapeutic management of primary central nervous system lymphoma in immunocompetent patients: Results of a critical review of the literature, <u>Ann. Oncol.</u> 8: 227-234 (1997)				
	EEEEER	Sandor <i>et al.</i> , Phase II trial of chemotherapy alone for primary CNS and intraocular lymphoma, <u>J. Clin. Oncol.</u> 16: 3000-3006 (1998)				
	FFFFR	Schabet et al., Diagnose und therapie der meningosis neoplastica, <u>Nervenarzt</u> 63: 317-27 (1992)				F
6	GGGGGR	Stamenkovic <i>et al.</i> , A B-lymphocyte activation molecule related to the nerve growth factor receptor and induced by cytokines in carcinomas, <u>EMBO J.</u> 8: 1403-1410 (1989)				
	ннннн	Stone <i>et al.</i> , A phase I study of bolus versus continuous infusion of the anti-CD19 immunotoxin, IgG-HD37-dgA, in patients with B-cell lymphoma, <u>Blood</u> 88: 1188-97 (1996)				
	IIIIR	Strauchen <i>et al.</i> , Chemotherapy in the management of intraocular lymphoma, Cancer 63: 1918-21 (1989)				
	JJJJJR	Tutt et al., Monoclonal antibody therapy of B cell lymphoma: Signaling activity on tumor cells appears more important than recruitment of effectors, <u>J. Immunol.</u> 161: 3176-3185 (1998)				
	KKKKKR	Valentine et al., B3.9 Structure and function of the B-cell specific 35-37 kDa CD20 protein, In: Leukocyte Typing III (McMichael, Ed., p. 440, Oxford University Press (1987)			··	
	LLLLLR	Valle et al., Activation of human B lymphocytes through CD40 and interleukin 4, Eur. J. Immunol. 19: 1463-1467 (1989)				
	MMMMMR	Valluri et al., Combination treatment of intraocular lymphoma, Retina 15: 125-9 (1995)				

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GN	NNNNR	van Besie 1991., Risk factors, treatment, and outcome of central nervous system recurrence in adults with intermediate-grade and immunistatic lymphoma, Blood 91: 1178-1184 (1998)	o o	'	CEN	父
THE	00000R	Verhoeyen et al., Reshaping human antibodies: Grafting an antilysozyme activity, Science 239: 1534-1536 (1988)	CENZ	0.	2003	S.
SEP 2 5 2000	👸	Vlasveld <i>et al.</i> , Treatment of low-grade non-Hodgkin's lymphoma with continuous infusion of low-dose recombinant interleukin-2 in combination with the B-cell-specific monoclonal antibody CLB-CD19, <i>Cancer Immunol. Immunother</i> . 40:37-47(1995)		160	200	
TE TRADENA	QQQQQR	Wang <i>et al.</i> , Induction of bcl-x by CD40 engagement rescues slg-induced apoptosis in murine B cells, <u>J. Immunol.</u> 155: 3722-5 (1995)				
	RRRRRR	White et al., Anti-CD20 monoclonal antibodies as novel treatments for non-Hodgkin's lymphoma, Pharm. Sci. Technol. Today 2: 95-101 (1999)				
-	SSSSSR	Wu et al., Pharmacokinetics and blood-brain barrier transport of an anti-transferrin receptor monoclonal antibody (OX26) in rats after chronic treatment with the antibody, <u>Drug. Metabol. Dispos.</u> 26: 937-9 (1998)			_	
	TTTTTR	Youle, Immunotoxins for central nervous system malignancy, <u>Semin. Cancer Biol.</u> 7: 65-70 (1996)				
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